



01/20/04

SIU 7399  
PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of Campbell

Serial No. 10/694,432

Filed October 27, 2003

For THERAPEUTIC USE OF D-METHIONINE TO REDUCE THE TOXICITY OF  
RADIATION

January 16, 2004

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SIR:

**INFORMATION DISCLOSURE STATEMENT**

\* In accordance with 37 C.F.R. 1.97 and 1.98 and MPEP 609, and in compliance with the  
duty of disclosure set forth in 37 C.F.R. 1.56, applicant submits the attached PTO/SB/08A for  
consideration by the Patent and Trademark Office in the above-entitled application and to be  
made of record therein.

The references listed on the attached PTO/SB/08A have been previously submitted in  
parent application Serial No. 09/911,195 and, therefore, in accordance with MPEP 609, are not  
being provided at this time.

This Information Disclosure Statement is being submitted pursuant to 37 C.F.R. §1.97(b)  
in that applicant believes that it is being filed prior to the mailing date of the first Office action  
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Disclosure Statement, the Commissioner is hereby authorized to charge any required fees  
regarding this Information Disclosure Statement to Deposit Account No. 19-1345.

Respectfully submitted,

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PTO/SB/08A  <b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>  <div style="display: flex; align-items: center;"> <div>           Use as many sheets as necessary)             1 of 8         </div> </div>		<b>Complete if Known</b>	
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		First Named Inventor	Campbell
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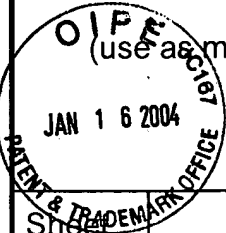
U.S. PATENT DOCUMENTS							
Examiner Initials*	Cite No. <sup>1</sup>	U.S. Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY		
		Number	Kind Code <sup>2</sup> (if known)				
	1	2,732,400		Weiss	01-24-1956		
	2	4,426,372		Borch	01-17-1984		
	3	5,002,755		Mitchell et al.	03-26-1991		
	4	5,292,773		Hirsch et al.	03-08-1994		
	5	5,430,064		Hirsch et al.	07-04-1995		
	6	5,466,678		Kawabata et al.	11-14-1995		
	7	6,177,434		Kopke et al.	01-23-2001		
FOREIGN PATENT DOCUMENTS							
Examiner Initials*	Cite No. <sup>1</sup>	Foreign Patent Document			Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	T <sup>6</sup>
		Office	Number <sup>4</sup>	Kind Code <sup>2</sup> (if known)			
	8	EP	06 200 04	A1	Fuji Kagaku Kogyo Kabushiki Kaisha	10-19-1994	

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OTHER ART - NON PATENT LITERATURE DOCUMENTS			
Examiner Initials*	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.) date, page(s), volume-issue number(s), publisher, city and/or country where published.	T <sup>6</sup>
	9	ALDEN, W. WESLEY et al., "Exacerbation of Cisplatin-Induced Nephrotoxicity by Methionine," <u>Chem.-Biol. Interactions</u> , 1984, 48, Pgs. 121-124	
	10	BASINGER, MARK A. et al., "Dithiocarbamate-Induced Biliary Platinum Excretion and the Control of cis-Platinum Nephrotoxicity," <u>Toxicology and Applied Pharmacology</u> , 1989, 97, Pgs. 279-288	
	11	BASINGER, MARK A. et al., "L-Methionine Antagonism of cis-Platinum Nephrotoxicity," <u>Toxicology and Applied Pharmacology</u> , 1990, 108, Pgs. 1-15	
	12	BASINGER, MARK A. et al., "L-Methionine Suppresses Pathological Sequelae of cis-Platinum in the Rat," <u>Fundamental and Applied Toxicology</u> , 1990, 14, Pgs. 568-577	
	13	BOOGAARD, PIETER J. et al., "4-Methylthiobenzoic Acid Reduces Cisplatin Nephrotoxicity In Rats Without Compromising Anti-Tumor Activity," <u>Biochemical Pharmacology</u> , 1991, Vol. 41, No. 12, Pgs. 1997-2003	
	14	BOOGAARD, PIETER J. et al., "The Role of Methallothionein in the Reduction of Cisplatin-Induced Nephrotoxicity by Bi <sup>3+</sup> -Pretreatment in the Rat <i>In Vivo</i> and <i>In Vitro</i> , Are Antioxidant Properties of Methallothionein More Relevant than Platinum Binding?", <u>Biochemical Pharmacology</u> , 1991, Vol. 41, No. 3, Pgs. 369-375	
	15	BURCHENAL, JOSEPH H. et al., "Studies of Cross-Resistance, Synergistic Combinations and Blocking of Activity of Platinum Derivates," <u>Biochimie</u> , 1978, 60, No. 9, Pgs. 961-965	
	16	CAMPBELL, KATHLEEN C.M. et al., "A Review of Cisplatin Protective Agents Emphasizing Nephro- and Otoprotectants," <u>Proposed Review Article Not Yet Submitted for Publication, Including Additional Reference Lists</u>	
	17	CAMPBELL, KATHLEEN C.M. et al., "D-Methionine Provides Excellent Protection from Cisplatin Ototoxicity in the Rat," <u>Hearing Research</u> , 1996, 102, Pgs. 90-98	

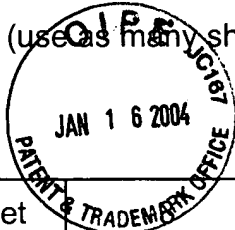
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	18	CAMPBELL, KATHLEEN C.M. et al., "D-Methionine Provides Protection Against Cisplatin Damage the Rat Stria Vascularis: A Semi-Quantitative Analysis," <u>Abstracts of the Twenty-First Annual Mid-Winter Research Meeting of the Association for Research in Otolaryngology</u> , February 15-19, 1998, Abstract No. 537, Pgs. 135	
	19	CARDINI, G. et al., "La Radioprotezione Dei Cromosomi Delle Cellule Midollari Umane In Vitro," <u>Radiobiologia Radioterapia E Fisica Medica</u> , 1967, Vol. 22, No. 6, Pgs. 371-375	
	20	CARRITHERS, S. L. et al., "Methylation Of Radiation Protector Compounds By Thiol Methyltransferase," <u>FASEB</u> , 1991, Vol. 5, No. 4, Pg. A824	
	21	CHURCH, MICHAEL W. et al., "The Comparative Effects of Sodium Thiosulfate, Diethyldithiocarbamate, Fosfomycin and WR-2721 on Ameliorating Cisplatin-Induced Ototoxicity," <u>Hearing Research</u> , 1995, 86/1,2, Pgs. 195-203	
	22	CORREA, J. N. et al., "Radiosensitization and radioprotection on murine chondrosarcoma," <u>Radiation Research</u> , 1978, Vol. 74, No. 3, Pg. 517	
	23	DE VECCHI, A., "Sperimentazione Clinica Di Una Nuova Sostanza Radioprotettiva (Cloruro Di Metil-Metionin-Sulfonio)," <u>Radiobiologia Radioterapia E Fisica Medica</u> , 1967, Vol. 22, No. 5, Pgs. 355-370	
	24	DEDON, PETER C. et al., "Characterization of the Reactions of Platinum Antitumor Agents with Biologic and Nonbiologic Sulfur-Containing Nucleophiles," <u>Biochemical Pharmacology</u> , 1987, Vol. 36, No. 12, Pgs. 1955-1964	
	25	DEEGAN, PATRICIA M. et al., "The nephrotoxicity, cytotoxicity and renal handling of a cisplatin-methionine complex in male Wistar rats," <u>Toxicology</u> , 1994, 89, Pgs. 1-14	
	26	DREWINKO, B. et al., "The Effect of <i>cis</i> -Diamminedichloroplatinum(II) on Cultured Human Lymphoma Cells and Its Therapeutic Implications," <u>Cancer Research</u> , December 1973, 33, Pgs. 3091-3095	
	27	DROBNIK, JAROSLAV et al., "Inactivation Of Bacteriophages With <i>Cis</i> -Platinum(II) Diamminedichloride," <u>Chem.-Biol. Interactions</u> , 1975, Pgs. 365-375	
	28	FRIEDMAN, M. E. et al., "The Blocking Of The Tetrachloroplatinate(II) Inhibition Of Malate Dehydrogenase By Sulfur-Containing Amino Acids," <u>Biochimica et Biophysica Acta</u> , 1974, 341, Pgs. 277-283	

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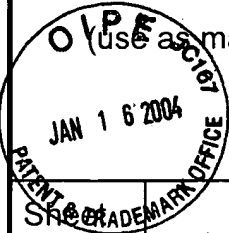
29	FRIEDMAN, MENDEL et al., "The Utilization and Safety of Isomeric Sulfur-Containing Amino Acids in Mice," <u>J. Nutr.</u> , 1984, 114, Pgs. 2301-2310	
30	GABAIZADEH, RAMIN et al., "Protection of Both Auditory Hair Cells and Auditory Neurons from Cisplatin Induced Damage," <u>Acta Otolaryngol</u> (Stockholm), 1997, 117, Pgs. 232-238	
31	GANDARA, DAVID R. et al., "Cisplatin Chemoprotection and Rescue: Pharmacologic Modulation of Toxicity," <u>Seminars in Oncology</u> , February 1991, Vol. 18, No. 1, Suppl. 3, Pgs. 49-55	
32	GANDARA, DAVID R. et al., "Evaluation of Cisplatin Dose Intensity: Current Status and Future Prospects," <u>Anticancer Research</u> , 1989, 9, Pgs. 1121-1128	
33	GESSLER, N. et al., "Antiradiation effects of S-methylmethionine (Vitamin U)," <u>Prikl. Biokhim. Mikrobiol.</u> , 1996, Vol. 32, No. 6, Pgs. 666-668	
34	GLOVER, DONNA et al., "Clinical Trials of WR-2721 and Cis-Platinum," <u>I. J. Radiation Oncology, Biology, Physics</u> , May 1989, Vol. 16, No. 5, Pgs. 1201-1204	
35	HANNERMANN, JORG et al., "Cisplatin-Induced Lipid Peroxidation and Decrease of Gluconeogenesis in Rat Kidney Cortex: Different Effects of Antioxidants and Radical Scavengers," <u>Toxicology</u> , 1988, 51, Pgs. 119-132	
36	HAYES, D. et al., "Amelioration Of Renal Toxicity Of High Dose Cis-Platinum Diammine Dichloride (CPDD) By Mannitol Induced Diuresis," <u>Proc. Am. Assoc. Cancer Res.</u> , 1976	
37	INFANTE, G. A. et al., "Chemical radioprotection on biological important compounds," <u>Radiation Research</u> , 1976, Vol. 67, No. 3, Pg. 637	
38	JONES, MARK M. et al., "Coadministration of Dimethyl Sulfoxide Reduces Cisplatin Nephrotoxicity," <u>Anticancer Research</u> , 1991, 11, Pgs. 1939-1942	
39	JONES, MARK M. et al., "Control of Nephrotoxicity in the Rat during Repeated cis-Platinum Treatments," <u>Journal of Applied Toxicology</u> , 1989, Vol. 9(4), Pgs. 229-233	
40	JONES, MARK M. et al., "Control of the Nephrotoxicity of Cisplatin by Clinically Used Sulfur-Containing Compounds," <u>Fundamental and Applied Toxicology</u> , 1992, 18, Pgs. 181-188	
41	JONES, MARK M. et al., "Inhibition of cis-diamminedichloroplatinum (II)-induced renal toxicity in the rat," <u>Cancer Chemotherapy and Pharmacology</u> , 1986, 17, Pgs. 38-42	

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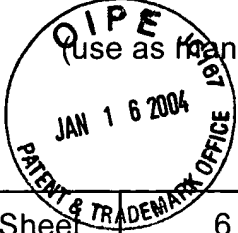
	42	JONES, MARK M. et al., "Relative effectiveness of some compounds for the control of cisplatin-induced nephrotoxicity," <u>Toxicology</u> , 1991, 68: Pgs. 227-247	
	43	JONES, MARK M. et al., "Thioether Suppression of Cisplatin Nephrotoxicity in the Rat," <u>Anticancer Research</u> , 1991, 11, Pgs. 449-454	
	44	JONES, MARK M. et al., "Thiol and Thioether Suppression of Cis-Platinum-Induced Nephrotoxicity in Rats Bearing the Walker 256 Carcinosarcoma," <u>Anticancer Research</u> , 1989, 9: 1937-1942	
	45	KIDO, K., "The influence of methylmethionine sulfonium chloride (MMS) on survivors of mice after X-ray irradiation, especially the consideration of the drug effect for the degeneration of intestinal mucosa," <u>Kansai Ika Daigaku Zasshi</u> , 1973, Vol. 25, No. 1, Pgs. 104-107	
	46	KIES, CONSTANCE et al., "Comparative Value of L-, DL-, and D-Methionine Supplementation of an Oat-based Diet for Humans," <u>J. Nutr.</u> , 1975, 105, Pgs. 809-814	
	47	KOPKE, RICHARD D. et al., "Use of Organotypic Cultures of Corti's Organ to Study the Protective Effects of Antioxidant Molecules on Cisplatin-Induced Damage of Auditory Hair Cells," <u>The American Journal of Otology</u> , 1997, 18, 559-571	
	48	KORVER, K. D. et al., "Round window application of D-methionine provides cisplatin otoprotection," <u>Abstracts of the Twenty-First Annual Mid-Winter Research Meeting of the Association for Research in Otolaryngology</u> , February 15-19, 1998, Abstract No. 536, Pg. 135	
	49	KOVACS, V. et al., "Study of the Radiation Protection Effect of Selenium-Methionine by Determining the Paramagnetic Properties of Liver Tissues of Mice," 1988, <u>Acta Physica Hungarica</u> , Vol. 64, No. 1-3, Pgs. 321-326	
	50	MEKHTIEV, M. A. et al., "Radioprotective effect during the separate and combined use of DL-methionine and thyroxine," <u>Database Chemabs Chemical Abstracts Service</u> , Abstract No. 76:54431, 1970 (see also <u>Tr. Inst. Fiziol., Akad. Nauk Azerb. SSR</u> , Vol. 11, pp. 83-100, 1970)	
	51	MELVIK, JAN EGIL et al., "Reduction of cis-Dichlorodiammineplatinum-Induced Cell Inactivation by Methionine," <u>Inorganica Chimica Acta</u> , 1987, 137, Pgs. 115-118	
	52	MERRIN, CLAUDE, "A New Method To Prevent Toxicity With High Doses Of Cis Diammine Platinum (Therapeutic Efficacy In Previously Treated Widespread And Recurrent Testicular Tumors)," <u>Proc. Am. Assoc. Cancer Res.</u> , 1976	

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53	MOLTENI, F. et al., "The use of S-adenosyl-methionine as a radioprotective agent," <u>Gazzetta Medica Italiana</u> , 1978, Vol. 137, No. 7-8, Pgs 303-308	
54	MONTINE, THOMAS J. et al., "Role of Endogenous Sulfur-Containing Nucleophiles in an In Vitro Model of cis-Diamminedichloroplatinum(II)-Induced Nephrotoxicity," <u>Biochemical Pharmacology</u> , 1990, Vol. 39, No. 11, Pgs. 1751-1757	
55	NAKANO, SACHIKO et al., "Potentiation of Cisplatin-Induced Lipid Peroxidation in Kidney Cortical Slices by Glutathione Depletion," <u>Japan. J. Pharmacol.</u> , 1989, 50, Pgs. 87-92	
56	ORMOND, T. et al., "Reduced Nephrotoxicity <u>In Vivo</u> and <u>In Vitro</u> of Cisplatin-Methionine Complex," <u>Brit. J. Pharmacol.</u> (suppl.), 1988, 95, Pg. 584P	
57	PRINTEN, KENNETH J. et al., "Utilization of D-methionine during total parenteral nutrition in postsurgical patients," <u>The American Journal of Clinical Nutrition</u> , June 1979, 32, Pgs. 1200-1205	
58	RAVI, RADHIKA P. et al., "Relationship of Pharmacodynamic Effects of Cisplatin to the Glutathione Levels in Cochlea, Inferior Colliculus and Kidney," <u>Pharmacologist</u> , 1991, 33(3), D-19, 402, P. 217	
59	RAVI, RADHIKA P. et al., "Diethyldithiocarbamate Protects Against Cisplatin Ototoxicity and Nephrotoxicity," <u>Otolaryngology Head and Neck Surgery</u> , 1992, 107(2), Poster 5, P. 232	
60	RESER, D. H. et al., "Physiological evidence for protection from cis-platin ototoxicity by D- and L-methionine in vivo," <u>Abstracts of the Twenty-First Annual Mid-Winter Research Meeting of the Association for Research in Otolaryngology</u> , February 15-19, 1998, Abstract No. 203, P. 51	
61	RHO, M. B. et al., "Structural evidence for protection from cisplatin ototoxicity by both D- and L-methionine in vivo," <u>Abstracts of the Twenty-First Annual Mid-Winter Research Meeting of the Association for Research in Otolaryngology</u> , February 15-19, 1998, Abstract No. 202, P. 51	
62	ROMITO, S., "Sulla radioprotezione cromosomica in vitro: esperienze con metionina, acido aspartico, leucina, lisina," <u>Fracastoro</u> , 1969, Vol. 62, No. 6, Pgs. 576-581	
63	SALIKHODZHAEV, Z. et al., "Stimulation of postirradiation recovery of rat haemopoiesis by a cobalt preparation," <u>Database Biosis Biosciences Information Service</u> , Philadelphia, Pennsylvania, Abstract No. 08095385 (see also <u>Radiobiologia</u> , Vol. 31, No. 6, pp. 835-837, 1991)	
64	SCHEIN, P. S., "Ethyol™ (WR-2721): a chemoprotective agent for platinum anti-cancer drugs," <u>Speaker Abstracts</u> (XP-002053095)	

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PTO/SB/08A		<b>Complete if Known</b>	
<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>  (use as many sheets as necessary)		Application Number	10/694,432
		Filing Date	October 27, 2003
		Confirmation Number	not yet assigned
		First Named Inventor	Campbell
		Group Art Unit	not yet assigned
Sheet 1 of 8	TRADEMARK OFFICE	Attorney Docket No.	SIU 7399

65	SCHWEITZER, VANESSA GAYL, "Cisplatin-Induced Ototoxicity: The Effect of Pigmentation and Inhibitory Agents," <u>Laryngoscope</u> , April 1993, 103, Pgs. 1-52
66	SHA et al., "Antioxidant therapy attenuates gentamicin-induced ototoxicity," <u>Abstracts of the Twenty-First Annual Mid-Winter Research Meeting of the Association for Research in Otolaryngology</u> , February 15-19, 1998, Abstract No. 535, P. 134
67	SPEER, R. J. et al., "Coordination Complexes of Platinum as Antitumor Agents," <u>Cancer Chemotherapy Reports</u> , 1975, Part I, Vol. 59, No. 3, Pgs. 629-641
68	SRINIVASAN, V. et al., "Radioprotection By Misoprostol (PGE, Methyl Analog) In Combination With Vitamin E, Selenomethionine and WR-3689794," <u>Eicosanoids and Other Bioactive Lipids in Cancer, Inflammation, and Radiation Injury 2</u> , edited by K. V. Honn et al., 1997, Plenum Press, New York, Pgs. 791-797
69	TOGNELLA, SERGIO, "Pharmacological interventions to reduce platinum-induced toxicity," <u>Cancer Treatment Reviews</u> , 1990, 17, Pgs. 139-142
70	TRESKES, MARCO et al., "WR2721 as a modulator of cisplatin- and carboplatin-induced side effects in comparison with other chemoprotective agents: a molecular approach," <u>Cancer Chemotherapy and Pharmacology</u> , 1993, 33, Pgs. 93-106
71	VAN DE WATER, T. et al., "Oxidative stress in the inner ear: Combinatorial therapy," (202) M. B. Rho et al., "Structural evidence for protection from cisplatin ototoxicity by both D- and L-methionine in vivo," <u>Abstracts of the Twenty-First Annual Mid-Winter Research Meeting of the Association for Research in Otolaryngology</u> , February 15-19, 1998, Abstract No. 6, P. 2
72	WALKER, ERNEST M., JR. et al., "Methods of Reduction of Cisplatin Nephrotoxicity," <u>Annals of Clinical and Laboratory Science</u> , 1981, Vol. 11, No. 5, Pgs. 397-409
73	WARD, J. M. et al., "Modification of the Renal Toxicity of <i>cis</i> -Dichlorodiammineplatinum(II) With Furosemide in Male F344 Rats," <u>Cancer Treatment Reports</u> , 1977, Vol. 61, No. 3, Pgs. 375-379
74	WHITWORTH, C. A. et al., "Alpha-lipoic acid as a protective agent against ototoxicity," (535) S. H. Sha et al., "Antioxidant therapy attenuates gentamicin-induced ototoxicity," <u>Abstracts of the Twenty-First Annual Mid-Winter Research Meeting of the Association for Research in Otolaryngology</u> , February 15-19, 1998, Abstract No. 532, P. 134

Examiner Signature	Date Considered
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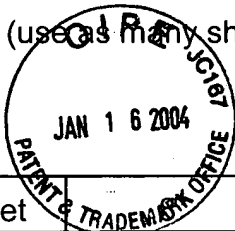
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	75	ZEZULKA, ALLISON YATES et al., "Nitrogen Retention in Men Fed Isolated Soybean Protein Supplemented with L-Methionine, D-Methionine, N-Acetyl-L-Methionine, or Inorganic Sulfate," <u>J. Nutr.</u> , 1976, 106, Pgs. 1286-1291	
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